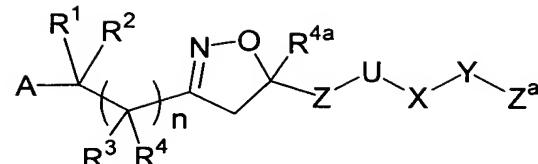


WHAT IS CLAIMED IS:

1. A compound of formula I or II:



I



II

5 or a stereoisomer or pharmaceutically acceptable salt form thereof, wherein;

A is $-\text{C}(\text{O})\text{NHOH}$, $-\text{C}(\text{O})\text{NHOR}^5$, $-\text{C}(\text{O})\text{NHOR}^6$, $-\text{N}(\text{OH})\text{COR}^5$, or $-\text{N}(\text{OH})\text{CHO}$;

U is absent or is O, $\text{NR}^{\text{a}1}$, $\text{C}(\text{O})$, $\text{CR}^{\text{a}}(\text{OH})$, $\text{C}(\text{O})\text{O}$, $\text{OC}(\text{O})$, $\text{C}(\text{O})\text{NR}^{\text{a}1}$, $\text{NR}^{\text{a}1}\text{C}(\text{O})$, $\text{OC}(\text{O})\text{O}$, $\text{OC}(\text{O})\text{NR}^{\text{a}1}$, $\text{NR}^{\text{a}1}\text{C}(\text{O})\text{O}$, $\text{NR}^{\text{a}1}\text{C}(\text{O})\text{NR}^{\text{a}1}$, $\text{S}(\text{O})_p$, $\text{S}(\text{O})_p\text{NR}^{\text{a}1}$,

10 $\text{NR}^{\text{a}1}\text{S}(\text{O})_p$, or $\text{NR}^{\text{a}1}\text{SO}_2\text{NR}^{\text{a}1}$;

X is absent or is C_{1-10} alkylene, C_{2-10} alkenylene, or C_{2-10} alkynylene;

Y is absent or is O, $\text{NR}^{\text{a}1}$, $\text{S}(\text{O})_p$, or $\text{C}(\text{O})$;

Z is a C_{3-13} carbocycle substituted with 1-5 R^b , or a 5-14 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and $\text{S}(\text{O})_p$, and substituted with 1-5 R^b ;

15 Z^{a} is H, C_{3-13} carbocycle substituted with 1-5 R^c , or a 5-14 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and $\text{S}(\text{O})_p$, and substituted with 1-5 R^c ;

provided that U, Y, Z, and Z^{a} do not combine to form a N-N, N-O, O-N, O-O, $\text{S}(\text{O})_p$ -O, O-S(O)_p, or S(O)_p-S(O)_p group;

20 R^1 is Q, C_{1-6} alkylene-Q, C_{2-6} alkenylene-Q, C_{2-6} alkynylene-Q, $-(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_r\text{O}(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_s\text{-Q}$, $-(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_r\text{NR}^{\text{a}}(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_s\text{-Q}$, $-(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_r\text{C}(\text{O})(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_s\text{-Q}$, $-(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_r\text{C}(\text{O})\text{O}(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_s\text{-Q}$, $-(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_r\text{OC}(\text{O})(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_s\text{-Q}$, $-(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_r\text{C}(\text{O})\text{NR}^{\text{a}}\text{R}^{\text{a}1}$, $-(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_r\text{C}(\text{O})\text{NR}^{\text{a}}(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_s\text{-Q}$, $-(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_r\text{NR}^{\text{a}}\text{C}(\text{O})\text{O}(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_s\text{-Q}$, $-(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_r\text{NR}^{\text{a}}\text{C}(\text{O})\text{NR}^{\text{a}}(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_s\text{-Q}$,

25 $-(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_r\text{OC}(\text{O})\text{O}(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_s\text{-Q}$, $-(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_r\text{OC}(\text{O})\text{NR}^{\text{a}}(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_s\text{-Q}$, $-(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_r\text{NR}^{\text{a}}\text{C}(\text{O})\text{NR}^{\text{a}}(\text{CR}^{\text{a}}\text{R}^{\text{a}1})_s\text{-Q}$,

-(CR^aR^{a1})_rS(O)_p(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rSO₂NR^a(CR^aR^{a1})_s-Q,
-(CR^aR^{a1})_rNR^aSO₂(CR^aR^{a1})_s-Q, or -(CR^aR^{a1})_rNR^aSO₂NR^a(CR^aR^{a1})_s-Q;

Q is, independently at each occurrence, H, CHF₂, CH₂F, CF₃, a C₃₋₁₃ carbocycle substituted with 0-5 R^d, or a 5-14 membered heterocycle consisting of 5 carbon atoms and 1-4 heteroatoms selected from N, O, and S(O)_p, and substituted with 0-5 R^d;

R² is Q¹, C₁₋₆ alkylene-Q¹, C₂₋₆ alkenylene-Q¹, C₂₋₆ alkynylene-Q¹,
-(CR^aR^{a1})_rO(CR^aR^{a1})_s-Q¹, -(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q¹,
-(CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q¹, -(CR^aR^{a1})_rC(O)O(CR^aR^{a1})_s-Q¹,
10 -(CR^aR^{a1})_rC(O)NR^aR^{a1}, or -(CR^aR^{a1})_rC(O)NR^a(CR^aR^{a1})_s-Q¹;
Q¹ is, independently at each occurrence, H, a C₃₋₁₃ carbocycle substituted with 0-5 R^d, or a 5-14 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and S(O)_p, and substituted with 0-5 R^d;

alternatively, R¹ and R² combine, along with the carbon atom to which they 15 are attached, to form a 3-7 membered carbocyclic or heterocyclic ring consisting of: carbon atoms and 0-3 heteroatoms selected from the group consisting of N, O, and S(O)_p and substituted with 0-2 R^d;

R³ is Q, C₁₋₆ alkylene-Q, C₂₋₆ alkenylene-Q, C₂₋₆ alkynylene-Q,
-(CR^aR^{a1})_rO(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q,
20 -(CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rC(O)O(CR^aR^{a1})_s-Q,
-(CR^aR^{a1})_rOC(O)(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rC(O)NR^aR^{a1},
-(CR^aR^{a1})_rC(O)NR^a(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^aC(O)(CR^aR^{a1})_s-Q,
-(CR^aR^{a1})_rOC(O)O(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rOC(O)NR^a(CR^aR^{a1})_s-Q,
25 -(CR^aR^{a1})_rNR^aC(O)O(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^aC(O)NR^a(CR^aR^{a1})_s-Q,
-(CR^aR^{a1})_rS(O)_p(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rSO₂NR^a(CR^aR^{a1})_s-Q,
-(CR^aR^{a1})_rNR^aSO₂(CR^aR^{a1})_s-Q, or -(CR^aR^{a1})_rNR^aSO₂NR^a(CR^aR^{a1})_s-Q;

alternatively, R¹ and R³ combine, along with the carbon atom to which they are attached, to form a 3-7 membered carbocyclic or heterocyclic ring consisting of:

carbon atoms and 0-3 heteroatoms selected from the group consisting of N, O, and S(O)_p and substituted with 0-2 R^d;

alternatively, when R¹ and R³ combine to form a carbocyclic or heterocyclic ring, the R² and R⁴ combine to form a double bond;

5 R⁴ is Q¹, C₁₋₆ alkylene-Q¹, C₂₋₆ alkenylene-Q¹, C₂₋₆ alkynylene-Q¹,
 -(CR^aR^{a1})_rO(CR^aR^{a1})_s-Q¹, -(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q¹,
 -(CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q¹, -(CR^aR^{a1})_rC(O)O(CR^aR^{a1})_s-Q¹,
 -(CR^aR^{a1})_rC(O)NR^aR^{a1}, or -(CR^aR^{a1})_rC(O)NR^a(CR^aR^{a1})_s-Q¹;

10 alternatively, R³ and R⁴ combine, along with the carbon atom to which they are attached, to form a 3-7 membered carbocyclic or heterocyclic ring consisting of: carbon atoms and 0-3 heteroatoms selected from the group consisting of N, O, and S(O)_p and substituted with 0-2 R^d;

15 R^{4a} is Q, C₁₋₆ alkylene-Q, C₂₋₆ alkenylene-Q, C₂₋₆ alkynylene-Q,
 -(CR^aR^{a1})_rO(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q,
 -(CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rC(O)O(CR^aR^{a1})_s-Q,
 -(CR^aR^{a1})_rC(O)NR^aR^{a1}, -(CR^aR^{a1})_rC(O)NR^aOR^a, -(CR^aR^{a1})_rC(O)NR^a(CR^aR^{a1})_s-Q,
 -(CR^aR^{a1})_rNR^aC(O)(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^aC(O)O(CR^aR^{a1})_s-Q,
 -(CR^aR^{a1})_rS(O)_p(CR^aR^{a1})_s-Q, or -(CR^aR^{a1})_rSO₂NR^a(CR^aR^{a1})_s-Q;

20 alternatively, R¹ and R^{4a} in Formula I combine, along with the carbon atoms to which they are attached, to form a 3-7 membered carbocyclic or heterocyclic ring consisting of: carbon atoms and 0-3 heteroatoms selected from the group consisting of N, O, and S(O)_p and substituted with 0-3 R^d, provided that n is 0;

25 alternatively, R³ and R^{4a} in Formula I combine, along with the carbon atoms to which they are attached, to form a 3-7 membered carbocyclic or heterocyclic ring consisting of: carbon atoms and 0-3 heteroatoms selected from the group consisting of N, O, and S(O)_p and substituted with 0-2 R^d;

R^a is, independently at each occurrence, H, C₁₋₆ alkyl, phenyl, or benzyl;

30 R^{a1} is, independently at each occurrence, H, C₁₋₆ alkyl substituted with 0-1 R^{c1}, C₂₋₆ alkenyl substituted with 0-1 R^{c1}, C₂₋₆ alkynyl substituted with 0-1 R^{c1}, or -(CH₂)_r-3-8 membered carbocyclic or heterocyclic ring consisting of carbon atoms

and 0-2 ring heteroatoms selected from N, NR^{a2}, O, and S(O)_p, and substituted with 0-3 R^{c1};

alternatively, R^a and R^{a1} when attached to a nitrogen, together with the nitrogen to which they are attached, combine to form a 5 or 6 membered heterocycle 5 consisting of carbon atoms and from 0-1 additional heteroatoms selected from N, NR^{a2}, O, and S(O)_p;

R^{a2} is, independently at each occurrence, C₁₋₄ alkyl, phenyl, or benzyl;

R^{a3} is, independently at each occurrence, H, C₁₋₆ alkyl substituted with 0-1 R^{c1}, C₂₋₆ alkenyl substituted with 0-1 R^{c1}, C₂₋₆ alkynyl substituted with 0-1 R^{c1}, or 10 -(CH₂)_r-3-8 membered carbocyclic or heterocyclic ring consisting of carbon atoms and 0-2 ring heteroatoms selected from N, NR^{a2}, O, and S(O)_p, and substituted with 0-3 R^{c1};

R^b is, independently at each occurrence, H, C₁₋₆ alkyl substituted with 0-1 R^{c1}, OR^a, SR^a, Cl, F, Br, I, =O, CN, NO₂, -NR^aR^{a1}, -C(O)R^a, -C(O)OR^a, 15 -C(O)NR^aR^{a1}, -C(S)NR^aR^{a1}, -NR^aC(O)NR^aR^{a1}, -OC(O)NR^aR^{a1}, -NR^aC(O)OR^a, -S(O)₂NR^aR^{a1}, -NR^aS(O)₂R^{a3}, -NR^aS(O)₂NR^aR^{a1}, -OS(O)₂NR^aR^{a1}, -S(O)_pR^{a3}, CF₃, CF₂CF₃, CHF₂, CH₂F, or phenyl;

R^c is, independently at each occurrence, H, OR^a, Cl, F, Br, I, =O, CN, NO₂, CF₃, CF₂CF₃, CH₂F, CHF₂, -(CR^aR^{a1})_rNR^aR^{a1}, -(CR^aR^{a1})_rC(=NCN)NR^aR^{a1}, 20 -(CR^aR^{a1})_rC(=NR^a)NR^aR^{a1}, -(CR^aR^{a1})_rC(=NOR^a)NR^aR^{a1}, -(CR^aR^{a1})_rC(O)NR^aOH, -(CR^aR^{a1})_rC(O)R^{a1}, -(CR^aR^{a1})_rC(O)OR^{a1}, -(CR^aR^{a1})_rC(S)OR^{a1}, -(CR^aR^{a1})_rC(O)NR^aR^{a1}, -(CR^aR^{a1})_rNR^aC(O)R^{a1}, -(CR^aR^{a1})_rC(S)NR^aR^{a1}, -(CR^aR^{a1})_rOC(O)NR^aR^{a1}, -(CR^aR^{a1})_rNR^aC(O)OR^{a1}, -(CR^aR^{a1})_rNR^aC(O)NR^aR^{a1}, 25 -(CR^aR^{a1})_rS(O)_pR^{a3}, -(CR^aR^{a1})_rSO₂NR^aR^{a1}, -(CR^aR^{a1})_rNR^aSO₂R^{a3}, -(CR^aR^{a1})_rNR^aSO₂NR^aR^{a1}, C₁₋₆ alkyl substituted with 0-2 R^{c1}, C₂₋₆ alkenyl substituted with 0-2 R^{c1}, C₂₋₆ alkynyl substituted with 0-2 R^{c1}, -(CR^aR^{a1})_r-C₃₋₁₀ carbocycle substituted with 0-2 R^{c1}, or -(CR^aR^{a1})_r-5-14 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and S(O)_p, and substituted with 0-2 R^{c1};

alternatively, when two R^c groups are attached to the same carbon atom, they form a 3-8 membered carbocyclic or heterocyclic spiro ring C substituted with 0-2 R^{c1} and consisting of carbon atoms, 0-4 ring heteroatoms selected from O, N, and S(O)_p, and 0-2 double bonds, provided that ring C contains other than a S-S, O-O, or 5 S-O bond;

alternatively, when two R^c groups are attached to adjacent carbon atoms, together with the carbon atoms to which they are attached they form a 5-7 membered carbocyclic or heterocyclic ring D substituted with 0-2 R^{c1} and consisting of: carbon atoms and 0-2 heteroatoms selected from N, O, and S(O)_p;

10 R^{c1} is, independently at each occurrence, H, C₁₋₄ alkyl, OR^a, Cl, F, Br, I, =O, CF₃, CN, NO₂, -C(O)R^a, -C(O)OR^a, -C(O)NR^aR^a, or -S(O)_pR^a;

R^d is, independently at each occurrence, C₁₋₆ alkyl, OR^a, Cl, F, Br, I, =O, CN, NO₂, -NR^aR^{a1}, -C(O)R^a, -C(O)OR^a, -C(O)NR^aR^{a1}, -C(O)NR^aOR^a, -C(S)NR^aR^{a1}, -NR^aC(O)NR^aR^{a1}, -OC(O)NR^aR^{a1}, -NR^aC(O)OR^a, -S(O)₂NR^aR^{a1}, -NR^aS(O)₂R^{a3}, 15 -NR^aS(O)₂NR^aR^{a1}, -OS(O)₂NR^aR^{a1}, -S(O)_pR^{a3}, CF₃, CF₂CF₃, C₃₋₁₀ carbocycle, or a 5-14 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and S(O)_p;

R⁵ is, independently at each occurrence, C₁₋₁₀ alkyl substituted with 0-2 R^b, or C₁₋₈ alkyl substituted with 0-2 R^e;

20 R^e is phenyl substituted with 0-2 R^b, or biphenyl substituted with 0-2 R^b;

R⁶ is, phenyl, naphthyl, C₁₋₁₀ alkyl-phenyl-C₁₋₆ alkyl-, C₃₋₁₁ cycloalkyl, C₁₋₆ alkylcarbonyloxy-C₁₋₃ alkyl-, C₁₋₆ alkoxy carbonyloxy-C₁₋₃ alkyl-, C₂₋₁₀ alkoxy carbonyl, C₃₋₆ cycloalkylcarbonyloxy-C₁₋₃ alkyl-, C₃₋₆ cycloalkoxycarbonyloxy-C₁₋₃ alkyl-, C₃₋₆ cycloalkoxycarbonyl,

25 phenoxycarbonyl, phenoxy carbonyloxy-C₁₋₃ alkyl-, phenylcarbonyloxy-C₁₋₃ alkyl-, C₁₋₆ alkoxy-C₁₋₆ alkylcarbonyloxy-C₁₋₃ alkyl-, [5-(C_{1-C}5 alkyl)-1,3-dioxa-cyclopenten-2-one-yl]methyl,

[5-(R^a)-1,3-dioxa-cyclopenten-2-one-yl]methyl,

(5-aryl-1,3-dioxa-cyclopenten-2-one-yl)methyl, -C₁₋₁₀ alkyl-NR⁷R^{7a},

30 -CH(R⁸)OC(=O)R⁹, or -CH(R⁸)OC(=O)OR⁹;

R⁷ is H, C₁₋₁₀ alkyl, C₂₋₆ alkenyl, C₃₋₆ cycloalkyl-C₁₋₃ alkyl-, or phenyl-C₁₋₆ alkyl-;

R^{7a} is H, C₁₋₁₀ alkyl, C₂₋₆ alkenyl, C₃₋₆ cycloalkyl-C₁₋₃ alkyl-, or phenyl-C₁₋₆ alkyl-;

5 R⁸ is H or C₁₋₄ linear alkyl;

R⁹ is H, C₁₋₈ alkyl substituted with 1-2 R^f, C₃₋₈ cycloalkyl substituted with 1-2 R^f, or phenyl substituted with 0-2 R^b;

R^f is, independently at each occurrence, C₁₋₄ alkyl, C₃₋₈ cycloalkyl, C₁₋₅ alkoxy, or phenyl substituted with 0-2 R^b;

10 n is 0 or 1;

p, at each occurrence, is selected from 0, 1, and 2;

r, at each occurrence, is selected from 0, 1, 2, 3, and 4; and

s, at each occurrence, is selected from 0, 1, 2, 3, and 4.

15 2. A compound according to Claim 1, wherein:

U is absent or is O, NR^{a1}, C(O), CR^a(OH), C(O)O, OC(O), C(O)NR^{a1}, NR^{a1}C(O), S(O)_p, S(O)_pNR^{a1}, or NR^{a1}S(O)_p;

X is absent or is C₁₋₃ alkylene or C₃₋₄ alkynylene;

Y is absent or is O, NR^{a1}, S(O)_p, or C(O);

20 Z is a C₅₋₁₀ carbocycle substituted with 1-3 R^b, or a 5-10 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and S(O)_p, and substituted with 1-3 R^b;

Z^a is H, C₃₋₁₃ carbocycle substituted with 1-3 R^c, or a 5-14 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and S(O)_p, and substituted with 1-3 R^c;

provided that U, Y, Z, and Z^a do not combine to form a N-N, N-O, O-N, O-O, S(O)_p-O, O-S(O)_p, or S(O)_p-S(O)_p group;

R¹ is Q, C₁₋₆ alkylene-Q, C₂₋₆ alkenylene-Q, C₂₋₆ alkynylene-Q, -(CR^aR^{a1})_rO(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q,

30 -(CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rC(O)O(CR^aR^{a1})_s-Q,

- (CR^aR^{a1})_rOC(O)(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rC(O)NR^aR^{a1},
- (CR^aR^{a1})_rC(O)NR^a(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^aC(O)(CR^aR^{a1})_s-Q,
- (CR^aR^{a1})_rOC(O)O(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rOC(O)NR^a(CR^aR^{a1})_s-Q,
- (CR^aR^{a1})_rNR^aC(O)O(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rS(O)_p(CR^aR^{a1})_s-Q,
- 5 -(CR^aR^{a1})_rSO₂NR^a(CR^aR^{a1})_s-Q, or -(CR^aR^{a1})_rNR^aSO₂(CR^aR^{a1})_s-Q;
- Q is, independently at each occurrence, H, CH₂F, CH₂Cl, CF₃, a C₃₋₁₃ carbocycle substituted with 0-3 R^d, or a 5-14 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and S(O)_p, and substituted with 0-3 R^d;
- 10 R² is Q¹, C₁₋₆ alkylene-Q¹, C₂₋₆ alkenylene-Q¹, C₂₋₆ alkynylene-Q¹,
- (CR^aR^{a1})_rO(CR^aR^{a1})_s-Q¹, -(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q¹,
- (CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q¹, -(CR^aR^{a1})_rC(O)O(CR^aR^{a1})_s-Q¹, or
- (CR^aR^{a1})_rC(O)NR^a(CR^aR^{a1})_s-Q¹;
- Q¹ is, independently at each occurrence, H, a C₃₋₁₀ carbocycle substituted with 0-3 R^d, or a 5-10 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and S(O)_p, and substituted with 0-3 R^d;
- 15 alternatively, R¹ and R², when attached to the same carbon atom, combine to form a 5-6 membered carbocyclic or heterocyclic ring consisting of: carbon atoms and 0-3 heteroatoms selected from the group consisting of N, O, and S(O)_p and substituted with 0-2 R^d;
- 20 R³ is Q, C₁₋₆ alkylene-Q, C₂₋₆ alkenylene-Q, C₂₋₆ alkynylene-Q,
- (CR^aR^{a1})_rO(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q,
- (CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rC(O)O(CR^aR^{a1})_s-Q,
- (CR^aR^{a1})_rC(O)NR^aR^{a1}, -(CR^aR^{a1})_rC(O)NR^a(CR^aR^{a1})_s-Q,
- 25 -(CR^aR^{a1})_rNR^aC(O)(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rOC(O)NR^a(CR^aR^{a1})_s-Q,
- (CR^aR^{a1})_rNR^aC(O)O(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^aC(O)NR^a(CR^aR^{a1})_s-Q,
- (CR^aR^{a1})_rS(O)_p(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rSO₂NR^a(CR^aR^{a1})_s-Q,
- (CR^aR^{a1})_rNR^aSO₂(CR^aR^{a1})_s-Q, or -(CR^aR^{a1})_rNR^aSO₂NR^a(CR^aR^{a1})_s-Q;

alternatively, R¹ and R³ combine, along with the carbon atom to which they are attached, to form a 4-7 membered carbocyclic or heterocyclic ring consisting of: carbon atoms and 0-3 heteroatoms selected from the group consisting of N, O, and S(O)_p and substituted with 0-2 R^d;

5 alternatively, when R¹ and R³ combine to form a carbocyclic or heterocyclic ring, the R² and R⁴ combine to form a double bond;

R⁴ is Q¹, C₁₋₆ alkylene-Q¹, C₂₋₆ alkenylene-Q¹, C₂₋₆ alkynylene-Q¹, -(CR^aR^{a1})_rO(CR^aR^{a1})_s-Q¹, -(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q¹, -(CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q¹, -(CR^aR^{a1})_rC(O)O(CR^aR^{a1})_s-Q¹, or

10 -(CR^aR^{a1})_rC(O)NR^a(CR^aR^{a1})_s-Q¹;

alternatively, R³ and R⁴ combine, along with the carbon atom to which they are attached, to form a 4-7 membered carbocyclic or heterocyclic ring consisting of: carbon atoms and 0-3 heteroatoms selected from the group consisting of N, O, and S(O)_p and substituted with 0-2 R^d;

15 R^{4a} is Q, C₁₋₆ alkylene-Q, C₂₋₆ alkenylene-Q, C₂₋₆ alkynylene-Q, -(CR^aR^{a1})_rO(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rC(O)O(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rC(O)NR^aR^{a1}, -(CR^aR^{a1})_rC(O)NR^aOR^a, -(CR^aR^{a1})_rC(O)NR^a(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^aC(O)(CR^aR^{a1})_s-Q, or -(CR^aR^{a1})_rSO₂NR^a(CR^aR^{a1})_s-Q;

20 alternatively, R¹ and R^{4a} in Formula I combine, along with the carbon atoms to which they are attached, to form a 4-7 membered carbocyclic or heterocyclic ring consisting of: carbon atoms and 0-3 heteroatoms selected from the group consisting of N, O, and S(O)_p and substituted with 0-3 R^d, provided that n is 0;

25 alternatively, R³ and R^{4a} in Formula I combine, along with the carbon atoms to which they are attached, to form a 4-7 membered carbocyclic or heterocyclic ring consisting of: carbon atoms and 0-3 heteroatoms selected from the group consisting of N, O, and S(O)_p and substituted with 0-2 R^d;

R^a is, independently at each occurrence, H, C₁₋₆ alkyl, phenyl, or benzyl;

R^{a1} is, independently at each occurrence, H, C₁₋₆ alkyl, C₂₋₆ alkenyl, C₂₋₆ alkynyl, or -(CH₂)_r-3-8 membered carbocyclic or heterocyclic ring consisting of carbon atoms and 0-2 ring heteroatoms selected from N, NR^{a2}, O, and S(O)_p;

alternatively, R^a and R^{a1} when attached to a nitrogen, together with the 5 nitrogen to which they are attached, combine to form a 5 or 6 membered heterocycle consisting of carbon atoms and from 0-1 additional heteroatoms selected from N, NR^{a2}, O, and S(O)_p;

R^c is, independently at each occurrence, H, OR^a, Cl, F, Br, =O, CN, NO₂, CF₃, CH₂F, CHF₂, CF₂CF₃, -(CR^aR^{a1})_rNR^aR^{a1}, -(CR^aR^{a1})_rC(O)R^{a1}, -(CR^aR^{a1})_rC(O)OR^{a1}, -(CR^aR^{a1})_rC(O)NR^aR^{a1}, -(CR^aR^{a1})_rNR^aC(O)R^{a1}, -(CR^aR^{a1})_rS(O)_pR^{a3}, -(CR^aR^{a1})_rSO₂NR^aR^{a1}, -(CR^aR^{a1})_rNR^aSO₂R^{a3}, C₁₋₆ alkyl substituted with 0-1 R^{c1}, C₂₋₆ alkenyl substituted with 0-1 R^{c1}, C₂₋₆ alkynyl substituted with 0-1 R^{c1}, -(CH₂)_r-C₃₋₆ carbocycle substituted with 0-2 R^{c1}, or -(CH₂)_r-5-6 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and S(O)_p, and substituted with 0-2 R^{c1};

alternatively, when two R^c groups are attached to the same carbon atom, they form a 3-8 membered carbocyclic or heterocyclic spiro ring C substituted with 0-2 R^{c1} and consisting of carbon atoms, 0-4 ring heteroatoms selected from O, N, and S(O)_p, and 0-2 double bonds, provided that ring C contains other than a S-S, O-O, or S-O bond;

alternatively, when two R^c groups are attached to adjacent carbon atoms, together with the carbon atoms to which they are attached they form a 5-7 membered carbocyclic or heterocyclic ring D substituted with 0-2 R^{c1} and consisting of: carbon atoms and 0-2 heteroatoms selected from N, O, and S(O)_p;

R^d is, independently at each occurrence, C₁₋₆ alkyl, OR^a, Cl, F, Br, =O, CN, NO₂, -NR^aR^{a1}, -C(O)R^a, -C(O)OR^a, -C(O)NR^aR^{a1}, -S(O)₂NR^aR^{a1}, -NR^aS(O)₂R^{a3}, -S(O)_pR^{a3}, CF₃, C₃₋₆ carbocycle, or a 5-6 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and S(O)_p;

R^5 is, independently at each occurrence, C₁₋₆ alkyl substituted with 0-2 R^b, or 30 C₁₋₄ alkyl substituted with 0-2 R^e; and

R^f is, independently at each occurrence, C_{1-4} alkyl, C_{3-6} cycloalkyl, C_{1-5} alkoxy, or phenyl substituted with 0-2 R^b .

3. A compound according to Claim 2, wherein:

5 A is $-C(O)NHOH$ or $-N(OH)CHO$;

U is absent or is O , NR^{a1} , $C(O)$, $CR^a(OH)$, $C(O)NR^{a1}$, $NR^{a1}C(O)$, $S(O)_p$,
 $S(O)_pNR^{a1}$, or $NR^{a1}S(O)_p$;

X is absent or is methylene, ethylene, propynylene, or butynylene;

Z is a C_{5-10} carbocycle substituted with 1-2 R^b , or a 5-10 membered

10 heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and
 $S(O)_p$, and substituted with 1-2 R^b ;

Z^a is H, C_{5-10} carbocycle substituted with 1-3 R^c , or a 5-10 membered
heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and
 $S(O)_p$, and substituted with 1-3 R^c ;

15 provided that U , Y , Z , and Z^a do not combine to form a N-N, N-O, O-N, O-O,
 $S(O)_p$ -O, O- $S(O)_p$, or $S(O)_p$ - $S(O)_p$ group;

R^1 is Q , C_{1-6} alkylene- Q , C_{2-6} alkenylene- Q , C_{2-6} alkynylene- Q ,
 $-(CR^aR^{a1})_rO(CR^aR^{a1})_s-Q$, $-(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q$,
 $-(CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q$, $-(CR^aR^{a1})_rC(O)O(CR^aR^{a1})_s-Q$,

20 $-(CR^aR^{a1})_rC(O)NR^aR^{a1}$, $-(CR^aR^{a1})_rC(O)NR^a(CR^aR^{a1})_s-Q$,
 $-(CR^aR^{a1})_rNR^aC(O)(CR^aR^{a1})_s-Q$, $-(CR^aR^{a1})_rOC(O)NR^a(CR^aR^{a1})_s-Q$,
 $-(CR^aR^{a1})_rNR^aC(O)O(CR^aR^{a1})_s-Q$, $-(CR^aR^{a1})_rS(O)_p(CR^aR^{a1})_s-Q$,
 $-(CR^aR^{a1})_rSO_2NR^a(CR^aR^{a1})_s-Q$, or $-(CR^aR^{a1})_rNR^aSO_2(CR^aR^{a1})_s-Q$;

Q is, independently at each occurrence, H, a C_{3-8} carbocycle substituted with
25 0-3 R^d , or a 5-10 membered heterocycle consisting of carbon atoms and 1-4
heteroatoms selected from N, O, and $S(O)_p$, and substituted with 0-3 R^d ;

R^2 is Q^1 , C_{1-6} alkylene- Q^1 , C_{2-6} alkenylene- Q^1 , $-CR^aR^{a1})_rO(CR^aR^{a1})_s-Q^1$,
 $-(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q^1$, or $-(CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q^1$;

Q^1 is, independently at each occurrence, H, a C₅₋₁₀ carbocycle substituted with 0-2 R^d, or a 5-10 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and S(O)_p, and substituted with 0-2 R^d;

R³ is Q, C₁₋₆ alkylene-Q, C₂₋₆ alkenylene-Q, C₂₋₆ alkynylene-Q,

5 -(CR^aR^{a1})_rO(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q,
 -(CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^aC(O)(CR^aR^{a1})_s-Q,
 -(CR^aR^{a1})_rNR^aC(O)O(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rNR^aC(O)NR^a(CR^aR^{a1})_s-Q,
 -(CR^aR^{a1})_rS(O)_p(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rSO₂NR^a(CR^aR^{a1})_s-Q, or
 -(CR^aR^{a1})_rNR^aSO₂(CR^aR^{a1})_s-Q;

10 R⁴ is Q¹, C₁₋₆ alkylene-Q¹, C₂₋₆ alkenylene-Q¹, -(CR^aR^{a1})_rO(CR^aR^{a1})_s-Q¹,
 -(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q¹, or -(CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q¹;
 R^{4a} is Q, C₁₋₄ alkylene-Q, -(CH₂)_rO(CH₂)_s-Q, -(CH₂)_rNR^a(CH₂)_s-Q,
 -(CH₂)_rC(O)(CH₂)_s-Q, -(CH₂)_rC(O)O(CH₂)_s-Q, -(CH₂)_rC(O)NR^aR^{a1},
 -(CH₂)_rC(O)NR^aOR^a, -(CH₂)_rC(O)NR^a(CH₂)_s-Q, -(CH₂)_rNR^aC(O)(CH₂)_s-Q, or
 15 -(CH₂)_rNR^aC(O)O(CH₂)_s-Q;

R^{a3} is, independently at each occurrence, H, C₁₋₆ alkyl, C₂₋₆ alkenyl, or
 -(CH₂)_r-3-8 membered carbocyclic or heterocyclic ring consisting of carbon atoms
 and 0-2 ring heteroatoms selected from N, NR^{a2}, O, and S(O)_p, and substituted with
 0-3 R^{c1};

20 R^c is, independently at each occurrence, H, OR^a, Cl, F, Br, =O, CF₃, CH₂F,
 CHF₂, -(CR^aR^{a1})_rNR^aR^{a1}, -(CR^aR^{a1})_rC(O)R^{a1}, -(CR^aR^{a1})_rC(O)OR^{a1},
 -(CR^aR^{a1})_rC(O)NR^aR^{a1}, -(CR^aR^{a1})_rNR^aC(O)R^{a1}, -(CR^aR^{a1})_rS(O)_pR^{a3},
 -(CR^aR^{a1})_rSO₂NR^aR^{a1}, -(CR^aR^{a1})_rNR^aSO₂R^{a3}, C₁₋₆ alkyl, C₂₋₆ alkenyl,
 C₂₋₆ alkynyl, C₃₋₆ cycloalkyl substituted with 0-1 R^{c1}, phenyl substituted with 0-2
 25 R^{c1}, or 5-6 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms
 selected from N, O, and S(O)_p, and substituted with 0-2 R^{c1};

 alternatively, when two R^c groups are attached to adjacent carbon atoms,
 together with the carbon atoms to which they are attached they form a 5-7 membered

carbocyclic or heterocyclic ring D substituted with 0-2 R^{c1} and consisting of: carbon atoms and 0-2 heteroatoms selected from N, O, and S(O)_p;

R^d is, independently at each occurrence, C₁₋₆ alkyl, OR^a, Cl, F, Br, =O, -NR^aR^{a1}, -C(O)R^a, -C(O)OR^a, -C(O)NR^aR^{a1}, -S(O)₂NR^aR^{a1}, -NR^aS(O)₂R^{a3}, 5 -S(O)_pR^{a3}, CF₃, or phenyl;

R⁵ is, independently at each occurrence, C₁₋₄ alkyl substituted with 0-2 R^b, or C₁₋₄ alkyl substituted with 0-2 R^e;

r, at each occurrence, is selected from 0, 1, 2, and 3; and

s, at each occurrence, is selected from 0, 1, 2, and 3.

10

4. A compound according to Claim 3, wherein:

A is -C(O)NHOH;

Z is phenyl substituted with 1-2 R^b, naphthyl substituted with 1-2 R^b, or pyridyl substituted with 1-2 R^b;

15

Z^a is phenyl substituted with 1-3 R^c, naphthyl substituted with 1-3 R^c, or a heterocycle substituted with 1-3 R^c and selected from furanyl, tetrahydrofuranyl, thiazolyl, oxazolyl, imidazolyl, isothiazolyl, isoxazolyl, 4,5-dihydro-isoxazolyl, thienyl, triazolyl, thiadiazolyl, oxadiazolyl, pyridyl, pyrimidinyl, piperazinyl, piperidinyl, pyranyl, pyrazinyl, pyrazolyl, pyridoimidazolyl, pyrrolidinyl, pyrrolyl, 20 indolyl, indolinyl, benzimidazolyl, benzothiazinyl, benzofuranyl, benzothiophenyl, benzoxazolyl, benzthiazolyl, benztriazolyl, benzisoxazolyl, benzisothiazolyl, quinolinyl, tetrahydroquinolinyl, isoquinolinyl, tetrahydro-isoquinolinyl, indazolyl, isobenzofuranyl, isoindazolyl, isoindolinyl, isoindolyl, methylenedioxypyhenyl, quinazolinyl, 1,1-dioxido-2,3-dihydro-4H-1,4-benzothiazin-4-yl, 1,1-dioxido-3,4- 25 dihydro-2H-1-benzothiopyran-4-yl, 3,4-dihydro-2H-chomen-4-yl, 2H-chomen-4-yl, and pyrazolo[1,5-a]pyridinyl;

provided that U, Y, Z, and Z^a do not combine to form a N-N, N-O, O-N, O-O, S(O)_p-O, O-S(O)_p, or S(O)_p-S(O)_p group;

R¹ is selected from Q, C₁₋₆ alkylene-Q, -(CR^aR^{a1})_rO(CR^aR^{a1})_s-Q, 30 -(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rC(O)(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rC(O)O(CR^aR^{a1})_s-Q, -(CR^aR^{a1})_rC(O)NR^aR^{a1},

$-(CR^aR^{a1})_rC(O)NR^a(CR^aR^{a1})_s-Q$, $-(CR^aR^{a1})_rNR^aC(O)(CR^aR^{a1})_s-Q$, or
 $-(CR^aR^{a1})_rNR^aC(O)O(CR^aR^{a1})_s-Q$;

5 Q is, independently at each occurrence, H, a C_{3-6} carbocycle substituted with 0-3 R^d , or a 5-10 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and $S(O)_p$, and substituted with 0-3 R^d ;

R^2 is Q^1 or C_{1-6} alkylene- Q^1 ;

Q^1 is, independently at each occurrence, H, phenyl substituted with 0-2 R^d , or a 5-6 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and $S(O)_p$, and substituted with 0-2 R^d ;

10 R^3 is Q, C_{1-4} alkylene-Q, C_{2-4} alkenylene-Q, C_{2-4} alkynylene-Q,

$-(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q$, $-(CR^aR^{a1})_rNR^aC(O)(CR^aR^{a1})_s-Q$,
 $-(CR^aR^{a1})_rNR^aC(O)O(CR^aR^{a1})_s-Q$, $-(CR^aR^{a1})_rNR^aC(O)NR^a(CR^aR^{a1})_s-Q$,
 $-(CR^aR^{a1})_rS(O)_p(CR^aR^{a1})_s-Q$, $-(CR^aR^{a1})_rSO_2NR^a(CR^aR^{a1})_s-Q$, or
 $-(CR^aR^{a1})_rNR^aSO_2(CR^aR^{a1})_s-Q$;

15 R^4 is Q^1 or C_{1-6} alkylene- Q^1 ;

R^{4a} is Q, $-CH_2-Q$, $-CH_2O(CH_2)_s-Q$, $-CH_2NR^a(CH_2)_s-Q$, $-CH_2C(O)(CH_2)_s-Q$,
 $-CH_2C(O)O(CH_2)_s-Q$, $-CH_2C(O)NR^aR^{a1}$, $-(CH_2)_rC(O)NR^aOR^a$,
 $-CH_2C(O)NR^a(CH_2)_s-Q$, $-CH_2NR^aC(O)(CH_2)_s-Q$, or $-CH_2NR^aC(O)O(CH_2)_s-Q$;

R^{a1} is, independently at each occurrence, H, C_{1-6} alkyl, phenyl, or benzyl;

20 R^{a3} is, independently at each occurrence, H, C_{1-6} alkyl, phenyl, or benzyl;

R^c is, independently at each occurrence, H, OR^a , Cl, F, Br, =O, CF_3 , CH_2F ,
 CHF_2 , $-(CR^aR^{a1})_rNR^aR^{a1}$, $-(CR^aR^{a1})_rC(O)R^{a1}$, $-(CR^aR^{a1})_rC(O)OR^{a1}$,
 $-(CR^aR^{a1})_rC(O)NR^aR^{a1}$, $-(CR^aR^{a1})_rNR^aC(O)R^{a1}$, $-(CR^aR^{a1})_rS(O)_pR^{a3}$,
 $-(CR^aR^{a1})_rSO_2NR^aR^{a1}$, $-(CR^aR^{a1})_rNR^aSO_2R^{a3}$, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6}

25 alkynyl, phenyl substituted with 0-2 R^{c1} , or 5-6 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and $S(O)_p$, and substituted with 0-2 R^{c1} ; and

 alternatively, when two R^c groups are attached to adjacent carbon atoms, together with the carbon atoms to which they are attached they form a 5-6 membered

carbocyclic or heterocyclic ring D substituted with 0-2 R^{c1} and consisting of: carbon atoms and 0-2 heteroatoms selected from N, O, and S(O)_p.

5. A compound according to Claim 4, wherein:

5 U is absent or is O, NR^{a1}, C(O), CR^a(OH), C(O)NR^{a1}, or NR^{a1}C(O);
X is absent or is methylene or butynylene;
Y is absent or is O;
Z is phenyl substituted with 1-2 R^b;
Z^a is naphthyl substituted with 1-3 R^c, or a heterocycle substituted with 1-3 R^c
10 and selected from furanyl, tetrahydrofuranyl, thiazolyl, oxazolyl, imidazolyl, isothiazolyl, isoxazolyl, 4,5-dihydro-isoxazolyl, thienyl, triazolyl, thiadiazolyl, oxadiazolyl, pyridyl, pyrimidinyl, pyranyl, pyrazinyl, pyrazolyl, pyridoimidazolyl, pyrrolidinyl, pyrrolyl, quinolinyl, tetrahydroquinolinyl, isoquinolinyl, tetrahydro-isoquinolinyl, imidazolyl, benzimidazolyl, benzothiazinyl, benzofuranyl, benzothiophenyl, benzoxazolyl, benzthiazolyl, benztriazolyl, benzisoxazolyl, benzisothiazolyl, indolyl, indolinyl, indazolyl, isobenzofuranyl, isoindazolyl, isoindolinyl, isoindolyl, quinazolinyl, 1,1-dioxido-2,3-dihydro-4H-1,4-benzothiazin-4-yl, 1,1-dioxido-3,4-dihydro-2H-1-benzothiopyran-4-yl, 3,4-dihydro-2H-chomen-4-yl, 2H-chomen-4-yl, and pyrazolo[1,5-a]pyridinyl;

15 provided that U, Y, Z, and Z^a do not combine to form a N-N, N-O, O-N, O-O, S(O)_p-O, O-S(O)_p, or S(O)_p-S(O)_p group;

20 R¹ is Q, C₁₋₆ alkylene-Q, -(CR^aR^{a1})_rNR^a(CR^aR^{a1})_s-Q, or -(CR^aR^{a1})_rNR^aC(O)O(CR^aR^{a1})_s-Q;
Q is, independently at each occurrence, H, a C₃₋₆ carbocycle substituted with
25 0-3 R^d, or a 5-6 membered heterocycle consisting of carbon atoms and 1-4 heteroatoms selected from N, O, and S(O)_p, and substituted with 0-3 R^d;
R² is H or C₁₋₆ alkylene-Q¹;
R³ is Q, C₁₋₄ alkylene-Q, C₂₋₄ alkenylene-Q, C₂₋₄ alkynylene-Q, -(CH₂)_rNR^a(CH₂)_s-Q, -(CH₂)_rNR^aC(O)(CH₂)_s-Q, -(CH₂)_rNR^aC(O)O(CR^aR^{a1})_s-Q,
30 -(CH₂)_rNR^aC(O)NR^a(CH₂)_s-Q, -(CH₂)_rS(O)_p(CH₂)_s-Q, or -(CH₂)_rNR^aSO₂(CH₂)_s-Q;
R⁴ is H or C₁₋₆ alkylene-Q¹;

R^{4a} is Q, $-\text{CH}_2\text{Q}$, $-\text{CH}_2\text{O-Q}$, $-\text{CH}_2\text{NR}^a\text{Q}$, $-\text{CH}_2\text{C(O)}_s\text{Q}$, $-\text{CH}_2\text{C(O)}\text{O-Q}$,
 $-\text{CH}_2\text{C(O)}\text{NR}^a\text{R}^{a1}$, $-\text{C(O)}\text{NR}^a\text{OR}^a$, $-\text{CH}_2\text{C(O)}\text{NR}^a\text{Q}$, or $-\text{CH}_2\text{NR}^a\text{C(O)}\text{O-Q}$;
5 R^a is, independently at each occurrence, H, or C_{1-4} alkyl;
 R^{a1} is, independently at each occurrence, H, or C_{1-4} alkyl;
5 R^{a3} is, independently at each occurrence, H, C_{1-4} alkyl, phenyl, or benzyl;
 R^c is, independently at each occurrence, C_{1-6} alkyl, C_{2-6} alkenyl,
 C_{2-6} alkynyl, OR^a , Cl, F, Br, $=O$, CF_3 , CH_2F , CHF_2 , $-\text{NR}^a\text{R}^{a1}$, $-(\text{CR}^a\text{R}^{a1})_r\text{C(O)}\text{R}^{a1}$,
 $-(\text{CR}^a\text{R}^{a1})_r\text{C(O)}\text{OR}^{a1}$, $-(\text{CR}^a\text{R}^{a1})_r\text{C(O)}\text{NR}^a\text{R}^{a1}$, $-(\text{CR}^a\text{R}^{a1})_r\text{NR}^a\text{C(O)}\text{R}^{a1}$,
 $-(\text{CR}^a\text{R}^{a1})_r\text{S(O)}_p\text{R}^{a3}$, $-(\text{CR}^a\text{R}^{a1})_r\text{SO}_2\text{NR}^a\text{R}^{a1}$, $-(\text{CR}^a\text{R}^{a1})_r\text{NR}^a\text{SO}_2\text{R}^{a3}$, or phenyl; and
10 alternatively, when two R^c groups are attached to adjacent carbon atoms,
 together with the carbon atoms to which they are attached they form a 5-6 membered
 carbocyclic or heterocyclic ring D substituted with 0-1 R^{c1} and consisting of: carbon
 atoms and 0-2 heteroatoms selected from N, O, and $S(O)_p$.

15 **6.** A compound according to Claim 5, wherein:

U is absent or is O, NR^{a1} , $C(O)$, or $CR^a(OH)$;
 Y is absent;
 R^1 is H, C_{1-4} alkylene-Q, $-(\text{CH}_2)_r\text{NR}^a(\text{CH}_2)_s\text{Q}$, or
 $-(\text{CH}_2)_r\text{NR}^a\text{C(O)}\text{O}(\text{CR}^a\text{R}^{a1})_s\text{Q}$;
20 R^2 is H, CH_3 , CH_2CH_3 , $CH_2CH_2CH_3$, or $CH(CH_3)_2$;
 R^3 is Q, C_{1-4} alkylene-Q, $-(\text{CH}_2)_r\text{NR}^a(\text{CH}_2)_s\text{Q}$,
 $-(\text{CH}_2)_r\text{NR}^a\text{C(O)}(\text{CH}_2)_s\text{Q}$, $-(\text{CH}_2)_r\text{NR}^a\text{C(O)}\text{O}(\text{CR}^a\text{R}^{a1})_s\text{Q}$,
 $-(\text{CH}_2)_r\text{NR}^a\text{C(O)}\text{NR}^a(\text{CH}_2)_s\text{Q}$, $-(\text{CH}_2)_r\text{S}(\text{CH}_2)_s\text{Q}$, or $-(\text{CH}_2)_r\text{NR}^a\text{SO}_2(\text{CH}_2)_s\text{Q}$;
 R^4 is H, CH_3 , CH_2CH_3 , $CH_2CH_2CH_3$, or $CH(CH_3)_2$;
25 R^{4a} is Q, $-\text{CH}_2\text{Q}$, $-\text{CH}_2\text{O-Q}$, $-\text{CH}_2\text{NR}^a\text{Q}$, $-\text{CH}_2\text{C(O)}_s\text{Q}$, $-\text{CH}_2\text{C(O)}\text{O-Q}$,
 $-\text{CH}_2\text{C(O)}\text{NR}^a\text{R}^{a1}$, $-\text{C(O)}\text{NR}^a\text{OR}^a$, or $-\text{CH}_2\text{C(O)}\text{NR}^a\text{Q}$;
 r , at each occurrence, is selected from 0, 1, and 2; and
 s , at each occurrence, is selected from 0, 1, and 2.

7. A compound according to Claim 6, wherein:

U is O, NR^{a1}, or CR^a(OH);

Z^a is naphthyl substituted with 1-3 R^c, or a heterocycle substituted with 1-3 R^c and selected from pyridyl, quinolinyl, imidazolyl, benzimidazolyl, indolyl, 1,1-dioxido-2,3-dihydro-4H-1,4-benzothiazin-4-yl, 1,1-dioxido-3,4-dihydro-2H-1-benzothiopyran-4-yl, 3,4-dihydro-2H-chomen-4-yl, 2H-chomen-4-yl, pyrazolyl, and pyrazolo[1,5-a]pyridinyl;

10 R¹ is H, CH₃, CH₂CH₃, CH₂CH₂CH₃, CH(CH₃)₂, NH₂, or -NHC(O)OC(CH₃)₃;

R² is H or CH₃;

15 R³ is Q, C₁₋₄ alkylene-Q, -NR^a(CH₂)_s-Q, -NR^aC(O)(CH₂)_s-Q, -NR^aC(O)O(CR^aR^{a1})_s-Q, -NR^aC(O)NR^a(CH₂)_s-Q, -S(CH₂)_s-Q, or -NR^aSO₂(CH₂)_s-Q;

R^{4a} is Q, -CH₂-Q, -CH₂O-Q, -CH₂NR^a-Q, or -C(O)NR^aOR^a;

20 Q is, independently at each occurrence, H, phenyl substituted with 0-3 R^d, or a 5-6 membered heterocycle consisting of carbon atoms and 1-2 heteroatoms selected from N, O, and S(O)_p, and substituted with 0-3 R^d;

R^b is, independently at each occurrence, H, C₁₋₆ alkyl, OR^a, Cl, F, Br, -NR^aR^{a1}, -C(O)R^a, -C(O)OR^a, -C(O)NR^aR^{a1}, -S(O)₂NR^aR^{a1}, -NR^aS(O)₂R^{a3}, -S(O)_pR^{a3}, or CF₃;

25 R^c is, independently at each occurrence, C₁₋₆ alkyl, C₂₋₆ alkenyl, C₂₋₆ alkynyl, OR^a, Cl, F, Br, =O, CF₃, -NR^aR^{a1}, -(CR^aR^{a1})_rC(O)R^{a1}, -(CR^aR^{a1})_rC(O)OR^{a1}, -(CR^aR^{a1})_rC(O)NR^aR^{a1}, -(CR^aR^{a1})_rNR^aC(O)R^{a1}, -(CR^aR^{a1})_rS(O)_pR^{a3}, -(CR^aR^{a1})_rSO₂NR^aR^{a1}, or -(CR^aR^{a1})_rNR^aSO₂R^{a3}; and

alternatively, when two R^c groups are attached to adjacent carbon atoms, together with the carbon atoms to which they are attached they form a 5-6 membered saturated ring consisting of: carbon atoms and 0-1 heteroatoms selected from N, O, and S(O)_p.

8. A compound according to Claim 1, wherein the compound is selected from the group:

N-hydroxy-2-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-
isoxazol-5-yl}-acetamide;

5 N-hydroxy-2-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-
isoxazol-5-yl}-propionamide;

N-hydroxy-2-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-5-morpholin-4-
ylmethyl-4,5-dihydro-isoxazol-5-yl}-acetamide;

10 N-hydroxy-2-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-5-piperazin-1-
ylmethyl-4,5-dihydro-isoxazol-5-yl}-acetamide;

2-{5-dimethylaminomethyl-3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-
4,5-dihydro-isoxazol-5-yl}-N-hydroxy-acetamide;

N-hydroxy-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-
isoxazol-5-yl}-propionamide;

15 N-hydroxy-3-methyl-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-
dihydro-isoxazol-5-yl}-butyramide;

(1-hydroxycarbamoyl-2-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-
dihydro-isoxazol-5-yl}-ethyl)-carbamic acid tert-butyl ester;

20 2-amino-N-hydroxy-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-
dihydro-isoxazol-5-yl}-propionamide;

3-amino-N-hydroxy-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-
dihydro-isoxazol-5-yl}-propionamide;

N-hydroxy-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-
isoxazol-5-yl}-3-methylsulfanyl-propionamide;

25 N-hydroxy-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-
isoxazol-5-yl}-3-morpholin-4-yl-propionamide;

3-amino-N-hydroxy-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-
dihydro-isoxazol-5-yl}-butyramide;

30 furan-2-carboxylic acid (2-hydroxycarbamoyl-1-{3-[4-(2-methyl-quinolin-4-
ylmethoxy)-phenyl]-4,5-dihydro-isoxazol-5-yl}-ethyl)-amide;

N-hydroxy-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-
 isoxazol-5-yl}-3-pyrrolidin-1-yl-propionamide;
 3-acetyl-amino-N-hydroxy-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-
 4,5-dihydro-oxazol-5-yl}-propionamide;
 5 3-dimethylamino-N-hydroxy-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-
 phenyl]-4,5-dihydro-oxazol-5-yl}-propionamide;
 3-(3-ethyl-ureido)-N-hydroxy-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-
 phenyl]-4,5-dihydro-oxazol-5-yl}-propionamide;
 N-hydroxy-3-methanesulfonylamino-3-{3-[4-(2-methyl-quinolin-4-
 10 ylmethoxy)-phenyl]-4,5-dihydro-oxazol-5-yl}-propionamide;
 3-[(furan-2-ylmethyl)-amino]-N-hydroxy-3-{3-[4-(2-methyl-quinolin-4-
 ylmethoxy)-phenyl]-4,5-dihydro-oxazol-5-yl}-propionamide;
 3-benzylamino-N-hydroxy-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-
 4,5-dihydro-oxazol-5-yl}-propionamide;
 15 (2-hydroxycarbamoyl-1-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-
 dihydro-oxazol-5-yl}-ethyl)-carbamic acid isobutyl ester;
 N-hydroxy-3-{5-methyl-3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-
 dihydro-oxazol-5-yl}-propionamide;
 N-hydroxy-3-{5-hydroxymethyl-3-[4-(2-methyl-quinolin-4-ylmethoxy)-
 20 phenyl]-4,5-dihydro-oxazol-5-yl}-propionamide;
 5-(2-hydroxycarbamoyl-ethyl)-3-[4-(2-methyl-quinolin-4-ylmethoxy)-
 phenyl]-4,5-dihydro-oxazole-5-carboxylic acid methyl ester;
 5-(2-hydroxycarbamoyl-ethyl)-3-[4-(2-methyl-quinolin-4-ylmethoxy)-
 phenyl]-4,5-dihydro-oxazole-5-carboxylic acid hydroxyamide;
 25 2-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-oxazol-5-yl}-
 cyclopent-1-enecarboxylic acid hydroxyamide;
 cis-2-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-oxazol-5-
 yl}-cyclopentanecarboxylic acid hydroxyamide;
 cis-4-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-oxazol-5-
 30 yl}-pyrrolidine-3-carboxylic acid hydroxyamide;
 cis-4-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-oxazol-5-
 yl}-tetrahydro-furan-3-carboxylic acid hydroxyamide;

N-hydroxy-2-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-1-oxa-2-aza-spiro[4.5]dec-2-en-6-yl}-acetamide;

5 N-hydroxy-2-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-1,8-dioxa-2-aza-spiro[4.5]dec-2-en-6-yl}-acetamide;

6-hydroxycarbamoylmethyl-3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-1-oxa-2,8-diaza-spiro[4.5]dec-2-ene-8-carboxylic acid tert-butyl ester;

N-hydroxy-2-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-1-oxa-2,8-diaza-spiro[4.5]dec-2-en-6-yl}-acetamide;

10 N-hydroxy-2-{8-methyl-3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-1-oxa-2,8-diaza-spiro[4.5]dec-2-en-6-yl}-acetamide;

2-{8-acetyl-3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-1-oxa-2,8-diaza-spiro[4.5]dec-2-en-6-yl}-N-hydroxy-acetamide;

7-methyl-3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-6-oxo-1-oxa-2,7-diaza-spiro[4.4]non-2-ene-9-carboxylic acid hydroxyamide;

15 7-methyl-3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-6-oxo-1-oxa-2,7-diaza-spiro[4.5]dec-2-ene-10-carboxylic acid hydroxyamide;

N-hydroxy-2-(4-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-isoxazol-5-yl}-tetrahydro-pyran-4-yl)-acetamide;

20 2-(1-acetyl-4-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-isoxazol-5-yl}-piperidin-4-yl)-N-hydroxy-acetamide;

3-hydroxycarbamoylmethyl-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-isoxazol-5-yl}-pyrrolidine-1-carboxylic acid tert-butyl ester;

N-hydroxy-2-(3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-isoxazol-5-yl}-pyrrolidin-3-yl)-acetamide; and

25 N-hydroxy-2-(1-methyl-3-{3-[4-(2-methyl-quinolin-4-ylmethoxy)-phenyl]-4,5-dihydro-isoxazol-5-yl}-pyrrolidin-3-yl)-acetamide;

or a stereoisomer or pharmaceutically acceptable salt form thereof.

30 9. A pharmaceutical composition, comprising: a pharmaceutically acceptable carrier and a therapeutically effective amount of a compound according to Claim 1 or a pharmaceutically acceptable salt form thereof.

10. A method for treating an inflammatory disorder, comprising: administering to a patient in need thereof a therapeutically effective amount of a compound according to Claim 1 or a pharmaceutically acceptable salt form thereof.

5

11. A method of treating a condition or disease mediated by MMPs, TACE, aggrecanase, or a combination thereof in a mammal, comprising: administering to the mammal in need of such treatment a therapeutically effective amount of a compound according to Claim 1 or a pharmaceutically acceptable salt form thereof.

10
15 12. A method comprising: administering a compound according to Claim 1 or a pharmaceutically acceptable salt form thereof, in an amount effective to treat a condition or disease mediated by MMPs, TACE, aggrecanase, or a combination thereof.

13. A method of treating according to Claim 12, wherein the disease or condition is selected from to as acute infection, acute phase response, age related macular degeneration, alcoholic liver disease, allergy, allergic asthma, anorexia, aneurism, aortic aneurism, asthma, atherosclerosis, atopic dermatitis, autoimmune disease, autoimmune hepatitis, Bechet's disease, cachexia, calcium pyrophosphate dihydrate deposition disease, cardiovascular effects, chronic fatigue syndrome, chronic obstruction pulmonary disease, coagulation, congestive heart failure, corneal ulceration, Crohn's disease, enteropathic arthropathy, Felty's syndrome, fever, fibromyalgia syndrome, fibrotic disease, gingivitis, glucocorticoid withdrawal syndrome, gout, graft versus host disease, hemorrhage, HIV infection, hyperoxic alveolar injury, infectious arthritis, inflammation, intermittent hydrarthrosis, Lyme disease, meningitis, multiple sclerosis, myasthenia gravis, mycobacterial infection, neovascular glaucoma, osteoarthritis, pelvic inflammatory disease, periodontitis, polymyositis/dermatomyositis, post-ischaemic reperfusion injury, post-radiation asthenia, psoriasis, psoriatic arthritis, pulmonary emphysema, pydoderma gangrenosum, relapsing polychondritis, Reiter's syndrome, rheumatic fever,

rheumatoid arthritis, sarcoidosis, scleroderma, sepsis syndrome, Still's disease, shock, Sjogren's syndrome, skin inflammatory diseases, solid tumor growth and tumor invasion by secondary metastases, spondylitis, stroke, systemic lupus erythematosus, ulcerative colitis, uveitis, vasculitis, and Wegener's granulomatosis.

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14. A method for treating inflammatory disorders, comprising: administering, to a host in need of such treatment, a therapeutically effective amount of a compound of Claim 1 or a pharmaceutically acceptable salt form thereof, in combination with one or more additional anti-inflammatory agents selected from selective COX-2 inhibitors, interleukin-1 antagonists, dihydroorotate synthase inhibitors, p38 MAP kinase inhibitors, TNF- α inhibitors and TNF- α antibody or protein sequestration agents.

10

15. An article of manufacture, comprising:

15

(a) a first container;

(b) a pharmaceutical composition located within the first container, wherein the composition, comprises: a first therapeutic agent, comprising: a compound of Claim 1 or a pharmaceutically acceptable salt form thereof; and,

20 (c) a package insert stating that the pharmaceutical composition can be used for the treatment of an inflammatory disorder.

16. An article of manufacture according to Claim 18, further comprising:

25 (d) a second container;
wherein components (a) and (b) are located within the second container and component (c) is located within or outside of the second container.

17. An article of manufacture, comprising:

30 (a) a first container;
(b) a pharmaceutical composition located within the first container, wherein the composition, comprises: a first therapeutic agent, comprising: a compound of Claim 1, or a pharmaceutically acceptable salt form thereof; and,

(c) a package insert stating that the pharmaceutical composition can be used in combination with a second therapeutic agent to treat an inflammatory disorder.

18. An article of manufacture according to Claim 20, further comprising:

5 (d) a second container;

wherein components (a) and (b) are located within the second container and component (c) is located within or outside of the second container.